

### **Amendments to the Specification**

*Please replace the paragraph at page 16, lines 2-20 with the following replacement paragraph:*

(3) The tested compound having desired concentration was reacted in the solution of 20mM tris-HCl (pH 7.5), 1mM of  $MgCl_2$ , 100  $\mu M$  of EDTA, 330  $\mu g/mL$  of bovine serum albumin, 4 $\mu g/mL$  of 5'-nucleotidase, 0.1  $\mu Ci$  of  $^3H$ -cAMP (0.064  $\mu M$  of cAMP), 10  $\mu M$  of rolipram in storage solution of PDE 7 for 2 hours at 25°C. After the reaction, suspension of ~~Sephadex~~ SEPHADEX®-QAE (cross-linked dextran gel) in 10mM of HEPES-Na (pH 7.0) was added to the reaction mixture, and the mixture was left at rest for 5 minutes. Further, ~~Sephadex~~ SEPHADEX®-QAE (cross-linked dextran gel) was added to the obtained supernatant and the mixture was leaved at rest for 5 minutes, then, the radioactivity of the solution was measured.

*Please replace the paragraph at page 17, lines 11-20 with the following replacement paragraph:*

(1) The active fraction of PDE 4 (phosphodiesterase IV) was obtained. That is, the livers obtained from three Balb/c mice (male, 12 weeks: obtainable from CLEA Japan, Inc.) were suspended with 30mL of buffer solution B [20mM of bis-tris, 5mM of 2-mercaptoethanol, 2mM of benzamidine, 2mM of EDTA, 0.1mM of 4-(2-aminoethyl)benzensulfonyl hydrochloride, 50mM of sodium acetate; pH 6.5], then homogenized by ~~Polytron~~ POLYTRON® homogenizer. The homogenate were centrifuged under 25,000 × G for 10 minutes at 4°C. The supernatant was separated and thus obtained supernatant was further centrifuged under 100,000 × G for 60 minutes at 4°C, and then filtrated with 0.2  $\mu m$  filter to obtain the soluble fraction.

*Please replace the paragraph at page 17, line 30 to page 18, line 6 with the following replacement paragraph:*

(3) The tested compound having desired concentration was reacted in the solution of 20mM tris-HCl (pH 7.5), 1mM of  $MgCl_2$ , 100  $\mu M$  of EDTA, 330  $\mu g/mL$  of bovine serum albumin, 4  $\mu g/mL$  of 5'-nucleotidase, 0.1 $\mu Ci$  of  $^3H$ -cAMP (0.064  $\mu M$  of cAMP), and storage solution of PDE 4 for 2 hours at 25°C. After the reaction, suspension of ~~Sephadex~~ SEPHADEX®-QAE (cross-linked dextran gel) in 10mM of HEPES-Na (pH 7.0) was added to the reaction mixture, and the mixture was left at rest for 5 minutes. Further, ~~Sephadex~~ SEPHADEX®-QAE (cross-

linked dextran gel) was added to the obtained supernatant and the mixture was left at rest for 5 minutes, then, the radioactivity of the solution was measured.

*Please replace the paragraph at page 23, line 23 to page 16, line 1 with the following replacement paragraph:*

(1) The active fraction of PDE 7 (phosphodiesterase VII) was obtained. That is, MOLT-4 (obtainable from ATCC as ATCC No. CRL-1582), which was cell line of human acute lymphoblastic lymphoma T cells, was incubated in RPMI1640 culture medium containing 10% fetal bovine serum to obtain  $5 \times 10^8$  MOLT-4 cells. The cells were collected by centrifugation and suspended with 10mL of buffer solution A [25mM of tris-HCl, 5mM of 2-mercaptoethanol, 2mM of benzamidine, 2mM of EDTA, 0.1mM of 4-(2-aminoethyl)benzensulfonyl hydrochloride; pH 7.5], then homogenized by ~~Polytron~~ POLYTRON® homogenizer. The homogenate were centrifuged under  $25,000 \times G$  for 10 minutes at 4°C. The supernatant was separated and thus obtained supernatant was further centrifuged under  $100,000 \times G$  for 60 minutes at 4°C, and then filtrated with 0.2  $\mu m$  filter to obtain the soluble fraction.